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ring nodes :
    1 2 3 4 5 6
chain bonds :
    1-7 2-8 3-9 4-11 6-12 10-11 12-13 13-14 14-15 14-16 14-17
ring bonds :
    1-2 1-6 2-3 3-4 4-5 5-6
exact/norm bonds :
    1-2 1-6 1-7 2-3 2-8 3-4 3-9 4-5 5-6 13-14 14-15 14-16 14-17
exact bonds :
    4-11 6-12 10-11 12-13
Match level :
    1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS
Element Count :
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Node 17: Limited C,C1-5

L4

ANSWER 1 OF 1 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2001:112571 CAPLUS

DOCUMENT NUMBER:

134:326724

TITLE:

The controlled glycosylation of a protein with a

bivalent glycan: towards a new class of

glycoconjugates, glycodendriproteins

AUTHOR (S):

CORPORATE SOURCE:

Davis, Benjamin G. Department of Chemistry, University of Durham, Science

Laboratories, Durham, DH1 3LE, UK

SOURCE:

Chemical Communications (Cambridge, United Kingdom)

(2001), (4), 351-352

CODEN: CHCOFS; ISSN: 1359-7345 Royal Society of Chemistry

PUBLISHER: DOCUMENT TYPE:

Journal

LANGUAGE:

English

OTHER SOURCE(S):

CASREACT 134:326724

The use of a novel bivalent carbohydrate methanethiosulfonate modification reagent (I), based on a flexible, branched divalent core in a combined site-directed mutagenesis and chemical modification strategy has allowed the first controlled synthesis of a pure protein bearing a branched glycan or a first generation glycodendriprotein. Site-directed mutagenesis was used to introduce one Cys residue into the sequence of subtilisin Bacillus lentus (SBL) to produce variant SBL-S156C, which was reacted with I rapidly and quant. to give first-generation glycodendriprotein S156C-(S-a)2, which was purified and its structures confirmed by ES-MS anal.

336817-35-7

RL: RCT (Reactant); RACT (Reactant or reagent)

29

(preparation of glycoconjugates of cysteine-modified subtilisin as glycodendriproteins)

RN 336817-35-7 CAPLUS

D-glycero-D-manno-Octitol, 2,6-anhydro-7-deoxy-8-thio-, 8-methanesulfonate (9CI) (CA INDEX NAME)

Absolute stereochemistry.

REFERENCE COUNT:

THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

RN 219668-71-0 CAPLUS
CN β-D-Galactopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate
(9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-74-3 CAPLUS

CN β-D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl 4-O-(2,3,4,6-tetra-O-acetyl-β-D-galactopyranosyl)-, triacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

REFERENCE COUNT:

THERE ARE 81 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 10 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN

81

ACCESSION NUMBER:

2000:34887 CAPLUS

DOCUMENT NUMBER:

132:89792

TITLE:

Neoglycoproteins and their preparation by reacting cysteine-containing proteins mutants with glycosyl

thiosulfonate

INVENTOR(S):
PATENT ASSIGNEE(S):

Jones, J. Bryan; Davis, Benjamin G. Genencor International, Inc., USA

SOURCE:

PCT Int. Appl., 86 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE			APPLICATION NO.			DATE		
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WO 2000001712	A2 2000	0113	WO 19	99-US1513	8 1999	0702		
WO 2000001712	A3 2000	0511						
W: AL, AM,	AT, AU, AZ,	BA, BB,	BG, BR,	BY, CA,	CH, CN,	CU, CZ,	DE,	
DK, EE,	ES, FI, GB,	GD, GE,	GH, GM,	HR, HU,	ID, IL,	IN, IS,	JP,	
KE, KG,	KP, KR, KZ,	LC, LK,	LR, LS,	LT, LU,	LV, MD,	MG, MK,	MN,	
MW, MX,	NO, NZ, PL,	PT, RO,	RU, SD,	SE, SG,	SI, SK,	SL, TJ,	TM,	
TR, TT,	UA, UG, UZ,	VN, YU,	ZW, AM,	AZ, BY,	KG, KZ,	MD, RU,	TJ, TM	
RW: GH, GM,	KE, LS, MW,	SD, SL,	SZ, UG,	ZW, AT,	BE, CH,	CY, DE,	DK,	

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ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
              CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
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                                              AU 1999-52081
                        A1
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     AU 749383
                        B2
                                              EP 1999-937203
                                                                19990702
     EP 1093459
                        A2
                              20010425
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, SI, LT, LV, FI, RO
                                                                19990702
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     JP 2002519050
                                              US 2002-62970
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     US 2002146803
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                              20021010
                                           US 1998-91687P
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                                                                19980702
PRIORITY APPLN. INFO.:
                                           US 1999-131446P P 19990428
                                           US 1999-347029
                                                             A3 19990702
                                           WO 1999-US15138 W 19990702
     The present invention relates to a chemical modified mutant protein including
AB
     a cysteine residue substituted for a residue other than cysteine in a
     precursor protein, the substituted cysteine residue being subsequently
     modified by reacting the cysteine residue with a glycosylated
     thiosulfonate. Also, a method of producing the chemical modified mutant protein is provided. The present invention also relates to a glycosylated methanethiosulfonate. Another aspect of the present invention is a method
     of modifying the functional characteristics of a protein including
     providing a protein and reacting the protein with a glycosylated
     methanethiosulfonate reagent under conditions effective to produce a
     glycoprotein with altered functional characteristics as compared to the
     protein. In addition, the present invention relates to methods of determining the
     structure-function relationships of chemical modified mutant proteins. Thus,
     a number of glycosyl methanethiosulfonates were synthesized and reacted with
     the N62C, the S156C, the S166C, or the L217C mutants of Bacillus lentus
     subtilisin and the resulting neoglycoproteins were characterized. Thus,
     the L217C mutant was reacted with 2,3,4,6-tetra-O-acetyl-β-D-
     glucopyranosyl methanethiosulfonate to prepare a monoglucosylated enzyme
     containing 3 acetyl groups. This derivative had a kcat/KM that was 8-fold greater
     than that of the wild-type enzyme. Addnl. there was an improvement in
     specificity for ester vs. amide hydrolysis: the ratio
      (kcat/KM) esterase/(kcat/KM) amidase was 17.2-fold greater than that of the
                  This modified enzyme may find use in enzymic peptide
     wild-type.
     synthesis.
     219668-45-8P 219668-49-2P 219668-52-7P
     219668-55-0P 219668-58-3P 219668-62-9P
     219668-64-1P 219668-67-4P 219668-69-6P 219668-71-0P 219668-74-3P 254909-31-4P
     254909-32-5P 254909-33-6P 254909-34-7P
     254909-35-8P 254909-36-9P 254909-37-0P
     254909-38-1P 254909-39-2P 254909-40-5P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
      (Reactant or reagent)
         (neoglycoproteins and their preparation by reacting cysteine-containing proteins
        mutants with glycosyl thiosulfonate)
     219668-45-8 CAPLUS
     \beta-D-Glucopyranose, 1-thio-, 2,3,4,6-tetraacetate 1-methanesulfonate
```

Absolute stereochemistry. Rotation (-).

(9CI) (CA INDEX NAME)

CN

219668-49-2 CAPLUS RN α -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX CN

RN 219668-52-7 CAPLUS CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX

Absolute stereochemistry. Rotation (-).

RN 219668-55-0 CAPLUS

CN α-D-Mannopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-58-3 CAPLUS

CN β -D-Galactopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-62-9 CAPLUS

CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl 4-O- β -D-galactopyranosyl- (9CI) (CA INDEX NAME)

RN 219668-64-1 CAPLUS CN α -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-67-4 CAPLUS CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-69-6 CAPLUS CN α -D-Mannopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-71-0 CAPLUS CN β -D-Galactopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

RN 219668-74-3 CAPLUS CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl 4-0-(2,3,4,6-tetra-0-acetyl- β -D-galactopyranosyl)-, triacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 254909-31-4 CAPLUS CN β -D-Glucopyranose, 1-thio-, diacetate 1-methanesulfonate (9CI) (CA INDEX NAME)

CM 1

CRN 254909-30-3 CMF C7 H14 O7 S2

Absolute stereochemistry.

CM 2

CRN 64-19-7 CMF C2 H4 O2

RN 254909-32-5 CAPLUS
CN β-D-Glucopyranose, 1-thio-, triacetate 1-methanesulfonate (9CI) (CA INDEX NAME)

CM 1

CRN 254909-30-3 CMF C7 H14 O7 S2 Absolute stereochemistry.

CM 2

CRN 64-19-7 CMF C2 H4 O2

RN 254909-33-6 CAPLUS CN α -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl, diacetate (9CI) (CA INDEX NAME)

CM 1

CRN 219668-49-2 CMF C9 H18 O8 S2

Absolute stereochemistry. Rotation (+).

CM 2

CRN 64-19-7 CMF C2 H4 O2

RN 254909-34-7 CAPLUS CN α -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl, triacetate (9CI) (CA INDEX NAME)

CM 1

CRN 219668-49-2 CMF C9 H18 O8 S2

CRN 64-19-7 CMF C2 H4 O2

RN 254909-35-8 CAPLUS CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl, diacetate (9CI) (CA INDEX NAME)

CM 1

CRN 219668-52-7 CMF C9 H18 O8 S2

Absolute stereochemistry. Rotation (-).

CM 2

CRN 64-19-7 CMF C2 H4 O2

RN 254909-36-9 CAPLUS CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl, triacetate (9CI) (CA INDEX NAME)

CM :

CRN 219668-52-7 CMF C9 H18 O8 S2

CRN 64-19-7 CMF C2 H4 O2

RN 254909-37-0 CAPLUS CN α -D-Mannopyranoside, 2-[(methylsulfonyl)thio]ethyl, triacetate (9CI) (CA INDEX NAME)

CM :

CRN 219668-55-0 CMF C9 H18 O8 S2

Absolute stereochemistry. Rotation (+).

CM 2

CRN 64-19-7 CMF C2 H4 O2

RN 254909-38-1 CAPLUS CN β -D-Galactopyranoside, 2-[(methylsulfonyl)thio]ethyl, triacetate (9CI) (CA INDEX NAME)

CM 1

CRN 219668-58-3 CMF C9 H18 O8 S2

CRN 64-19-7 CMF C2 H4 O2

254909-39-2 CAPLUS RNCN

 β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl 4-O- β -D-galactopyranosyl-, pentaacetate (9CI) (CA INDEX NAME)

CM

219668-62-9 C15 H28 O13 S2 CRN CMF

Absolute stereochemistry. Rotation (+).

CM

CRN 64-19-7 CMF C2 H4 O2

254909-40-5 CAPLUS

β-D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl 4-O-β-D-galactopyranosyl-, hexaacetate (9CI) (CA INDEX NAME)

CM 1

CRN 219668-62-9 C15 H28 O13 S2 CMF

CRN 64-19-7 CMF C2 H4 O2

HO-C-CH3

ANSWER 11 OF 11 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

CORPORATE SOURCE:

1998:756618 CAPLUS

DOCUMENT NUMBER:

130:110474

TITLE:

Controlled Site-Selective Glycosylation of Proteins by

Pub 12/25/98

a Combined Site-Directed Mutagenesis and Chemical

Modification Approach

AUTHOR (S):

Davis, Benjamin G.; Lloyd, Richard C.; Jones, J. Bryan

Department of Chemistry, University of Toronto,

Toronto, ON, M5S 3H6, Can. SOURCE:

Journal of Organic Chemistry (1998), 63(26), 9614-9615

CODEN: JOCEAH; ISSN: 0022-3263

PUBLISHER:

American Chemical Society

DOCUMENT TYPE: LANGUAGE:

Journal English

Site-directed mutagenesis combined with chemical modification provides a general method that allows for both regio- and glycan-specific glycosylation of proteins. The strategy involves the introduction of cysteine at preselected positions and then reaction of its thiol residue with glycomethanethiosulfonate reagents. Four different sites of subtilisin Bacillus lentus (SBL) were mutated to cysteine (SBL-N62C, -S156C, -S166C, -L217C) and glycosylated using a series of protected and unprotected mono- and disaccharide methanethiosulfonates. Through adjustment of pH and appropriate selection of the glycosylation site, differently acetylated glycoforms of SBL were prepared

IT

CN

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent) (controlled site-selective glycosidation of proteins by a combined

site-directed mutagenesis and chemical modification approach)

219668-45-8 CAPLUS

 β -D-Glucopyranose, 1-thio-, 2,3,4,6-tetraacetate 1-methanesulfonate (9CI) (CA INDEX NAME)

219668-67-4P 219668-69-6P 219668-71-0P

219668-74-3P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent) (controlled site-selective glycosylation of proteins by a combined site-directed mutagenesis and chemical modification approach)

RN 219668-49-2 CAPLUS

CN α -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-52-7 CAPLUS

CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 219668-55-0 CAPLUS

CN α-D-Mannopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-58-3 CAPLUS

CN β-D-Galactopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX NAME)

RN 219668-62-9 CAPLUS

CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl 4-O- β -D-galactopyranosyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-64-1 CAPLUS

CN α -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-67-4 CAPLUS

CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-69-6 CAPLUS

CN α -D-Mannopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-71-0 CAPLUS

CN β -D-Galactopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

RN 219668-74-3 CAPLUS
CN β-D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl
4-0-(2,3,4,6-tetra-O-acetyl-β-D-galactopyranosyl)-, triacetate (9CI)
(CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

219668-45-8DP, reaction product with cysteine residues in IT mutagenized subtilisin 219668-49-2DP, reaction product with cysteine residues in mutagenized subtilisin 219668-52-7DP, reaction product with cysteine residues in mutagenized subtilisin 219668-55-0DP, reaction product with cysteine residues in mutagenized subtilisin 219668-58-3DP, reaction product with cysteine residues in mutagenized subtilisin 219668-62-9DP, reaction product with cysteine residues in mutagenized subtilisin 219668-64-1DP, reaction product with cysteine residues in mutagenized subtilisin 219668-67-4DP, reaction product with cysteine residues in mutagenized subtilisin 219668-69-6DP, reaction product with cysteine residues in mutagenized subtilisin 219668-71-0DP, reaction product with cysteine residues in mutagenized subtilisin 219668-74-3DP, reaction product with cysteine residues in mutagenized subtilisin RL: SPN (Synthetic preparation); PREP (Preparation) (controlled site-selective glycosylation of proteins by a combined site-directed mutagenesis and chemical modification approach) 219668-45-8 CAPLUS RN $\beta\text{-D-Glucopyranose, 1-thio-, 2,3,4,6-tetraacetate 1-methanesulfonate}$ CN (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 219668-49-2 CAPLUS CN α -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX NAME)

RN 219668-52-7 CAPLUS
CN β-D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 219668-55-0 CAPLUS
CN α-D-Mannopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-58-3 CAPLUS CN β -D-Galactopyranoside, 2-[(methylsulfonyl)thio]ethyl (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-62-9 CAPLUS CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl 4-O- β -D-galactopyranosyl- (9CI) (CA INDEX NAME)

RN 219668-64-1 CAPLUS CN α -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-67-4 CAPLUS CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-69-6 CAPLUS CN α -D-Mannopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 219668-71-0 CAPLUS CN β -D-Galactopyranoside, 2-[(methylsulfonyl)thio]ethyl, tetraacetate (9CI) (CA INDEX NAME)

RN 219668-74-3 CAPLUS CN β -D-Glucopyranoside, 2-[(methylsulfonyl)thio]ethyl 4-O-(2,3,4,6-tetra-O-acetyl- β -D-galactopyranosyl)-, triacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

REFERENCE COUNT:

THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT